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- (81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.
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For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.



(57) Abstract: The present invention provides a pharmaceutical composition and method for regenerating nerves and treating neurological diseases based on nerve regeneration. The present invention employs a substance, such as Pep5, PKC, IP3, Rho, Rho GDI, Rho kinase, or the like, which are involved in a p75 signal transduction pathway, or an agent capable of specifically interacting with any of these substances to block or suppress the p75 signal transduction pathway, thereby stopping inhibition of nerve regeneration. As a result, nerve regeneration is resumed. The present invention is also the first to disclose that the PTD domain is useful as an agent for nerve regeneration.





tional Application No

PCT/JP2004/004385 CLASSIFICATION OF SUBJECT MATTER PC 7 C07K7/08 A61k Ã6ĪK38/10 C12N15/11 G01N33/50 C12N15/62 A01K67/027 C12N15/10 A61P25/28 According to International Patent Classification (IPC) or to both national classification and IPC B. FIELDS SEARCHED Minimum documentation searched (classification system followed by classification symbols) IPC 7 C07K A61K C12N GO1N A01K A61P Documentation searched other than minimum documentation to the extent that such documents are included. In the fields searched Electronic data base consulted during the International search (name of data base and, where practical, search terms used) EPO-Internal, EMBASE, BIOSIS, WPI Data, PAJ, CHEM ABS Data, PASCAL C. DOCUMENTS CONSIDERED TO BE RELEVANT Category ° Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. P,X YAMASHITA TOSHIHIDE ET AL: "The p75 1-10.receptor acts as a displacement factor 41-48, that releases Rho from Rho-GDI." 50-58, NATURE NEUROSCIENCE. MAY 2003, 60,66, vol. 6, no. 5, May 2003 (2003-05), pages 71-75, 461-467, XP002295642 80,86, ISSN: 1097-6256 89, 91-93 97-115 the whole document -/--Further documents are listed in the continuation of box C. Patent family members are listed in annex. Special categories of cited documents: *T* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the "A" document defining the general state of the art which is not considered to be of particular relevance "E" earlier document but published on or after the international "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to filing date *L* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) involve an inventive step when the document is taken alone "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such docu-"O" document referring to an oral disclosure, use, exhibition or other means ments, such combination being obvious to a person skilled in the art. Po document published prior to the international filing date but later than the priority date claimed "&" document member of the same patent family Date of the actual completion of the International search Date of mailing of the international search report 10 September 2004 1 9. 01. 2005

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Ir ional Application No
PCT/JP2004/004385

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Y	abstract	1-10, 41-48, 50-58, 60,66, 71-75, 80,86, 89, 91-93, 97-115
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	In particular the last two paragraphs	97–115
	-/	
DOT TO A SOLE	(continuation of second sheet) (language 2004)	

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Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WONG SCOTT T ET AL: "A p75(NTR) and Nogo receptor complex mediates repulsive signaling by myelin-associated glycoprotein." NATURE NEUROSCIENCE. DEC 2002, vol. 5, no. 12, December 2002 (2002-12), pages 1302-1308, XP002295645 ISSN: 1097-6256 cited in the application abstract	1-10, 41-48, 50-58, 60,66, 71-75, 80,86, 89, 91-93, 97-115 61,66, 72,73, 75,80, 86,87, 89, 91-93,
		97,98, 100-102, 105-107
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	column 2, line 40 - line 56 column 5, line 9 - line 55 claims 1-10	105-107 1-10, 41-48, 50-58, 60,66, 71-75, 80,86, 89, 91-93, 97-115
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	page 1132, column 3, line 10 - line 12 page 1133; figure page 1133, column 3, paragraph 2 - paragraph 3	97–115
1	<u></u>	}

Ir Ional Application No

C.(Continua	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	FC1/JP2004/004385
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Υ		
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22 March 2001 (2001-03-22) page 8, line 30 - page 9, line 5 page 14, line 17 - page 15, line 15 example 1 claims 1-13 Y SCHWARZE S R ET AL: "PROTEIN TRANSDUCTION: UNRESTRICTED DELIVERY INTO ALL CELLS?" TRENDS IN CELL BIOLOGY, ELSEVIER SCIENCE LTD, XX, vol. 10, no. 7, July 2000 (2000-07), pages 290-295, XP001135090 ISSN: 0962-8924			Relevant to claim No.
TRANSDUCTION: UNRESTRICTED DELIVERY INTO ALL CELLS?" TRENDS IN CELL BIOLOGY, ELSEVIER SCIENCE LTD, XX, vol. 10, no. 7, July 2000 (2000-07), pages 290-295, XP001135090 ISSN: 0962-8924	Y	22 March 2001 (2001-03-22) page 8, line 30 - page 9, line 5 page 14, line 17 - page 15, line 15 example 1	41-48,
	Y	SCHWARZE S R ET AL: "PROTEIN TRANSDUCTION: UNRESTRICTED DELIVERY INTO ALL CELLS?" TRENDS IN CELL BIOLOGY, ELSEVIER SCIENCE LTD, XX, vol. 10, no. 7, July 2000 (2000-07), pages 290-295, XP001135090 ISSN: 0962-8924	41-48,

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Box II Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
Although claims 99-104, 108 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
2. Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box III Observations where unity of invention is lacking (Continuation of item 3 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
see additional sheet
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.: 1-10,41-48(part),50-58(part),60(part),61(part),66(part),72-74(part) 75(part),80(part),86-88(part),
Remark on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

INTERNATIONAL SEARCH REPORT Information on patent family members

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